

AMENDMENT TO THE CLAIMS:

This listing of claims will replace all prior listings of claims in the application:

LISTING OF CLAIMS:

1. (ORIGINAL) A MHC Class I-restricted epitope peptide derived from survivin, said epitope having at least one of the following characteristics:

(i) capable of binding to the Class I HLA molecule to which it is restricted at an affinity as measured by the amount of the peptide that is capable of half maximal recovery of the Class I HLA molecule (C_{50} value) which is at the most 50 μM as determined by the assembly binding assay as described herein,

(ii) capable of eliciting $\text{INF-}\gamma$ -producing cells in a PBL population of a cancer patient at a frequency of at least 1 per 10^4 PBLs as determined by an ELISPOT assay, and/or

(iii) capable of *in situ* detection in a tumor tissue of CTLs that are reactive with the epitope peptide.

2. (ORIGINAL) A peptide according to claim 1 having a C_{50} value, which is at the most 30 μM .

3. (ORIGINAL) A peptide according to claim 2 having a C_{50} value, which is at the most 20 μM .

4. (ORIGINAL) A peptide according to claim 1, which is restricted by a MHC Class I HLA-A molecule.

5. (ORIGINAL) A peptide according to claim 4, which is restricted by a MHC Class I HLA species selected from the group consisting of HLA-A1, HLA-A2, HLA-A3, HLA-A11 and HLA-A24.

6. (ORIGINAL) A peptide according to claim 5, which is restricted by HLA-A2.

7. (ORIGINAL) A peptide according to claim 6, which is selected from the group consisting of FLKLDRERA (SEQ ID NO:1), TLPPAWQPFL (SEQ ID NO:2), ELTLGEFLKL (SEQ ID NO:3), LLLGEFLKL (SEQ ID NO:4) and LMLGEFLKL (SEQ ID NO:5).

8. (ORIGINAL) A peptide according to claim 1, which is restricted by a MHC Class I HLA-B molecule.

9. (ORIGINAL) A peptide according to claim 8, which is restricted by a MHC Class I HLA-B species selected from the group consisting of HLA-B7, HLA -B35, HLA -B44, HLA-B8, HLA-B15, HLA-B27 and HLA-B51.

10. (ORIGINAL) A peptide according to claim 9, which is restricted by HLA-B35.

11. (ORIGINAL) A peptide according to claim 10, which is selected from the group consisting of CPTENEPDL (SEQ ID NO:6), EPDLAQCF (SEQ ID NO:7), CPTENEPDY (SEQ ID NO:8) and EPDLAQCFY (SEQ ID NO:9).

12. (ORIGINAL) A peptide according to claim 1 comprising at the most 20 amino acid residues.

13. (ORIGINAL) A peptide according to claim 12 that comprises at the most 10 amino acid residues.

14. (ORIGINAL) A peptide according to claim 1, which is a nonapeptide or a decapeptide.
15. (ORIGINAL) A peptide according to claim 1, which is a native sequence of survivin of a mammal species.
16. (ORIGINAL) A peptide according to claim 15 that is derived from human survivin.
17. (ORIGINAL) A peptide according to claim 1, which is derived from a native sequence of survivin by substituting, deleting or adding at least one amino acid residue.
18. (ORIGINAL) A peptide according to any of the preceding claims, which is phosphorylated.
19. (ORIGINAL) A peptide according to claim 18, which comprises Thr34 of the native survivin disclosed in US 6,45,23.
20. (ORIGINAL) A peptide according to claim 1 comprising, for each specific HLA allele, any of the amino acid residues as indicated in the following table:

HLA allele	Position 1	Position 2	Position 3	Position 5	Position 6	Position 7	C-terminal
HLA-A1		T,S	D,E			L	Y
HLA-A2		L, M			V		L, V
HLA-A3		L, V, M	F, Y				K, Y, F
HLA-A11		V, I, F, Y	M, L, F, Y , I				K, R
HLA-A23		I, Y					W, I
HLA-A24		Y		I, V	F		I, L, F
HLA-A25		M, A, T	I				W
HLA-A26	E, D	V, T, I, L, F			I, L, V		Y, F
HLA-A28	E, D	V, A, L					A, R
HLA-A29		E					Y, L
HLA-A30		Y, L, F, V					Y
HLA-A31			L, M, F, Y				R
HLA-A32		I, L					W
HLA-A33		Y, I, L, V					R
HLA-A34		V, L					R
HLA-A66	E, D	T, V					R, K
HLA-A68	E, D	T, V					R, K
HLA-A69		V, T, A					V, L
HLA-A74		T					V, L
HLA-B5		A, P	F, Y				I, L
HLA-B7		P					L, F
HLA-B8			K	K, R			L
HLA-B14		R, K					L, V
HLA-B15 (B62)		Q, L, K, P , H, V, I, M, S, T					F, Y, W
HLA-B17							L, V
HLA-B27		R					Y, K, F, L
HLA-B35		P					I, L, M, Y
HLA-B37		D, E					I, L, M
HLA-B38		H	D, E				F, L
HLA-B39		R, H					L, F
HLA-B40 (B60, 61)		E	F, I, V				L, V, A, W, M, T, R
HLA-B42		L, P					Y, L
HLA-B44		E					F, Y, W
HLA-B46		M, I, L, V					Y, F
HLA-B48		Q, K					L
HLA-B51		A, P, G					F, Y, I, V
HLA-B52		Q	F, Y				I, V

HLA-B53	P					W,F,L
HLA-B54	P					
HLA-B55	P					A,V
HLA-B56	P					A,V
HLA-B57	A,T,S					F,W,Y
HLA-B58	A,T,S					F,W,Y
HLA-B67	P					L
HLA-B73	R					P
HLA-Cw1	A,L					L
HLA-Cw2	A,L					F,Y
HLA-Cw3	A,L					L,M
HLA-Cw4	Y,P,F					L,M,F,Y
HLA-Cw6						L,I,V,Y
HLA-Cw6	Y					L,Y,F
HLA-Cw8	Y					L,I
HLA-Cw16	A,L					L,V

21. (ORIGINAL) A peptide according to claims 1 that is capable of eliciting INF- γ -producing cells in a PBL population of a cancer patient at a frequency of at least 10 per 104 PBLs.

22. (ORIGINAL) A peptide according to claim 1, which is capable of eliciting INF- γ -producing cells in a PBL population of a patient having a cancer disease where survivin is expressed.

23. (ORIGINAL) A peptide according to claim 22 where the cancer disease is selected from the group consisting of a haematopoietic malignancy including chronic lymphatic leukemia and chronic myeloid leukemia, melanoma, breast cancer, cervix cancer, ovary cancer, lung cancer, colon cancer, pancreas cancer and prostate cancer.

24. (ORIGINAL) A peptide according to claim 1, which is capable of eliciting INF- γ -producing cells in a PBL population of a patient having a cancer disease, said INF- γ -producing

cells having cytotoxic effect against survivin expressing cells of a cancer cell line, including a cell line selected from the group consisting of the breast cancer cell line MCF-7 and the melanoma cell line FM3.

25. (ORIGINAL) A pharmaceutical composition comprising the peptide according to claim 1.

26. (CURRENTLY AMENDED) A composition according to claim 25 that comprises a peptide having a sequence contained in native human survivin according to claim 4 in combination with another -a- modified survivin peptide that differs from a peptide which is contained in a native survivin by the presence of at least one substitution, deletion or addition amino acid modification according to claim 8.

27. (CURRENTLY AMENDED) A composition according to claim 26 comprising a native survivin peptide having the sequence contained in SEQ ID NO:36 (FTELTLGEF) in combination with another modified survivin peptide having the sequence contained in SEQ ID NO:14 (STFKNWPFL) according to claim 6 in combination with a peptide according to claim 10.

28. (ORIGINAL) A composition according to claim 25, which is a vaccine capable of eliciting an immune response against a cancer disease.

29. (ORIGINAL) A composition according to claim 25, further comprising an immunogenic protein or peptide fragment selected from a protein or peptide fragment not belonging to or derived from the survivin protein family.

30. (ORIGINAL) A composition according to claim 29, where the protein or peptide fragment not belonging to or derived from the survivin protein family is a protein, or peptide fragment hereof, involved in regulation of cell apoptosis.

31. (ORIGINAL) A composition according to claim 29 where the immunogenic protein or peptide fragment selected from a protein or peptide fragment hereof not belonging to or derived from the survivin protein family is Bcl-2 or a peptide fragment hereof.

32. (ORIGINAL) A composition according to claim 25, which is a multiepitope vaccine.

33. (ORIGINAL) A composition according to claim 28 where the vaccine is capable of eliciting an immune response against a cancer disease where survivin is expressed.

34. (ORIGINAL) A composition according to claim 33 where the cancer disease is selected from the group consisting of a haematopoietic malignancy including chronic lymphatic leukemia and chronic myeloid leukemia, melanoma, breast cancer, cervix cancer, ovary cancer, lung cancer, colon cancer, pancreas cancer and prostate cancer.

35. (ORIGINAL) A composition according to claim 33 or 34 where the vaccine elicits the production in the vaccinated subject of effector T-cells having a cytotoxic effect against the cancer cells.

36. (ORIGINAL) A composition for ex vivo or in situ diagnosis of the presence in a cancer patient of survivin reactive T-cells among PBLs or in tumor tissue, the composition comprising a peptide according to claim 1.

37. (ORIGINAL) A diagnostic kit for ex vivo or in situ diagnosis of the presence in a cancer patient of survivin reactive T-cells among PBLs or in tumour tissue comprising a peptide according to claim 1.

38. (ORIGINAL) A complex of a peptide according to claims 1 and a Class I HLA molecule or a fragment of such molecule.

39. (ORIGINAL) A complex according to claim 38 which is monomeric.

40. (ORIGINAL) A complex according to claim 38 which is multimeric.

41. (ORIGINAL) A method of detecting in a cancer patient the presence of survivin reactive T-cells, the method comprising contacting a tumour tissue or a blood sample with a complex according to claim 38 and detecting binding of the complex to the tissue or the blood cells.

42. (ORIGINAL) A molecule that is capable of binding specifically to a peptide according to claims 1.

43. (ORIGINAL) A molecule according to claim 36 which is an antibody or a fragment hereof.

44. (ORIGINAL) A molecule that is capable of blocking the binding of a molecule according to claim 42 or 43.

45. (ORIGINAL) A method of treating a cancer disease, the method comprising administering to a patient suffering from the disease an effective amount of the composition according to claim 25, a molecule according to claim 42 or a molecule according to claim 44.

46. (ORIGINAL) A method according to claim 45 wherein the disease to be treated is a cancer disease where survivin is expressed.

47. (ORIGINAL) A method according to claim 46 wherein the cancer disease is selected from the group consisting of a haematopoietic malignancy including chronic lymphatic leukemia and chronic myeloid leukemia, melanoma, breast cancer, cervix cancer, ovary cancer, lung cancer, colon cancer, pancreas cancer and prostate cancer.

48. (ORIGINAL) A method according to claim 45, which is combined with a further treatment.

49. (ORIGINAL) A method according to claim 48 wherein the further treatment is radiotherapy or chemotherapy.